



(11) Publication number : **0 419 261 B1**

(12)

EUROPEAN PATENT SPECIFICATION

(45) Date of publication of patent specification :
02.06.93 Bulletin 93/22

(51) Int. Cl.⁵ : **B65B 31/00**

(21) Application number : **90310306.7**

(22) Date of filing : **20.09.90**

(54) **Method of and apparatus for introducing into a container a suspension or solution of a material in a propellant held under pressure.**

(30) Priority : **21.09.89 GB 8921337**

(43) Date of publication of application :
27.03.91 Bulletin 91/13

(45) Publication of the grant of the patent :
02.06.93 Bulletin 93/22

(84) Designated Contracting States :
AT BE CH DE DK ES FR IT LI NL SE

(56) References cited :
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US-A- 3 642 035

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EP 0 419 261 B1

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Description

This invention relates to a method and apparatus for introducing into a container a pharmaceutical substance to be dispensed in aerosol form and a propellant therefor.

Conventionally, pharmaceutical substances which are to be dispensed in aerosol form are usually suspended in a mixture of at least two propellants, at least one of which has a high enough boiling point to be liquid at room temperature, and at least one of which has a low enough boiling point to be a gas at room temperature.

For convenience, these are referred to below as a liquid, or low pressure, propellant, and a gaseous, or high pressure, propellant respectively. The pharmaceutical material is first suspended in the liquid propellant by a mixing operation. Each aerosol container is then partly filled with this suspension. A quantity of the gaseous propellant is then introduced into each of the containers using either a cold-fill method or a high-pressure method. In the former, the filling operation is carried out at a temperature sufficiently below room temperature for the gaseous propellant to be liquid. Each container is then closed by a closure which includes an outlet valve through which the contents of the container can subsequently be dispensed. In the high-pressure method, the closure is applied to the container before the gaseous propellant is introduced, and that propellant is introduced subsequently into each container by forcing it under pressure into the container through the outlet valve, which during this operation acts in effect as an inlet valve.

No satisfactory method currently exists for filling a container with a suspension or solution of a pharmaceutical in a single or multi-component propellant which is gaseous at room temperature. It is an object of one aspect of the present invention to provide such a method, and to provide an apparatus for carrying out that method.

Figures 1 and 2 of the accompanying drawings show diagrammatically a typical known apparatus for introducing into a container a pharmaceutical material and a two-component propellant system.

Figure 1 shows the introduction into a container C of a suspension of a pharmaceutical material in a liquid propellant. Vessel 1 contains a bulk supply of this suspension which is pumped by a pump 2, through a non-return valve 3, into a metering cylinder 4 provided with a vent 5. From there, the suspension passes to a filling head 8. In the inoperative condition the suspension passes through the head 8 to a non-return valve 9 and thence back to the vessel. The suspension is thus kept constantly in circulation. When a quantity of suspension is to be introduced into the container C, the container is positioned below the head 8, and the valves 3 and 9 are closed. The pneu-

matic cylinder 6 is then operated to force the piston therein downwardly, thus increasing the pressure in the suspension trapped between the valves 3 and 9 to a level sufficiently to open a valve in filling head 8 and to cause suspension to pass from the filling head into the container C. The valves 3 and 9 are then opened and the valve in filling head 8 shut, and when the piston in the cylinder 6 is withdrawn to its original position the cylinder 4 refills from the vessel 1. Movement of the filling nozzle into and out of engagement with each can is effected by a piston and cylinder arrangement 7. The filling head 8 is arranged to operate only when it is in engagement with a container C.

The apparatus of Figure 1 introduces into container C a suspension of pharmaceutical material in a liquid propellant, and after an aerosol valve is crimped on the can C the apparatus shown in Figure 2 operates on it to introduce gaseous propellant. The apparatus of Figure 2 is formed of components which are substantially the same in principle as corresponding components of Figure 1, except that there is nothing corresponding to the non-return valve 9 and there is no recycling. Components in Figure 2 are denoted by reference numerals which correspond to those used in Figure 1, with the addition of 10. The vessel 11 contains gaseous propellant only, under sufficient pressure for it to be a liquid, and contains no pharmaceutical material.

When the apparatus of Figure 2 is in operation, a small quantity of gaseous propellant escapes each time the filling head 18 is lifted from a container C. This is of no particular consequence provided the amount of propellant lost in this way is small.

However, this feature of the operation of the apparatus of Figure 2 means that were it used for introducing into a container a suspension or solution of a pharmaceutical material in a high pressure propellant, it would be entirely unsatisfactory. It can be seen that if vessel 11 contained such a suspension or solution, what would escape each time the filling head 18 was lifted from a container would be a quantity of such a suspension or solution. This would present a hazard to workers involved in the operation, and where the pharmaceutical material concerned was an expensive one, could also represent a significant financial loss. Furthermore, the escaped pharmaceutical material would tend to deposit on the surrounding part of the apparatus and on the exterior of the container itself, giving rise to problems of cleaning. The first of these problems could be avoided, in theory, by surrounding the apparatus of Figure 2 by an exhaust system, though this would involve considerable expense. The other two problems would not be avoided even by such an exhaust system.

According to the present invention there is provided a method of introducing into a container a suspension or solution of a pharmaceutical substance in a propellant held under pressure, the suspension or

solution being circulated in a line which includes a filling head, which method comprises bringing the filling head into communication with the container; introducing a quantity of such suspension or solution into the container through the filling head; introducing a quantity of high pressure propellant without any of the said substance into the filling head while it is still in communication with the container, thereby to flush through any suspension or solution remaining in the filling head; and withdrawing the filling head from the container.

According to the present invention there is further provided an apparatus for introducing into a container a suspension or solution of a pharmaceutical substance in a propellant held under pressure, which comprises a line in which the suspension or solution circulates, the line including a filling head adapted to be brought into and out of communication with the container; means for supplying to the filling head along the line a quantity of the said suspension or solution; and means for supplying to the filling head a quantity of high pressure propellant without any of the said substance, the filling head being so arranged that the flow of propellant without any of the said substance flushes out any suspension or solution remaining in the filling head.

The invention also provides a filling head for use in introducing into a container a suspension or solution of a pharmaceutical substance in a propellant held under pressure, comprising an outlet adapted to communicate, in use, with the container; first and second inlets each communicating with the said outlet via a common flow path, through which inlets, in use, propellant containing the said substance, and high pressure propellant not containing the said substance, are respectively introduced; and means for selectively closing the first and second inlets so that fluid entering either inlet cannot flow out of the other.

In all aspects of the invention, it is advantageous and convenient if the propellant without the suspension or solution is the same propellant as that in which the substance is held.

Advantageously, the propellant is 1,1,1,2-tetrafluoroethane (also known as propellant "134a").

Preferably, the pharmaceutical substance being filled into the container is salbutamol or beclomethasone dipropionate.

Preferred embodiments of the invention are described below, by example only, with reference to Figures 3, 4A-4D and 5 of the accompanying drawings, in which:

Figure 3 shows diagrammatically the apparatus of the present invention;

Figures 4A-4D show one embodiment of a filling head which may be used in the apparatus of Figure 3; and

Figure 5 shows another embodiment of a filling head which may be used in the apparatus of Fig-

ure 3.

The apparatus according to the invention shown in Figure 3 comprises, in effect, something resembling a combination of the apparatus of Figures 1 and 2, but with a common filling head of a novel design. The components shown in Figure 3 are denoted by reference numerals which correspond to those shown in Figure 1, but with the addition of 20 or 30. The vessel 21 contains a suspension of a pharmaceutical substance in a high-pressure propellant, and the vessel 31 contains a supply of the same propellant alone, i.e. without any pharmaceutical substance suspended therein. Although the vessel 31 here contains the same propellant, a different high pressure propellant can of course be used. Furthermore, vessel 21 might contain a solution of the pharmaceutical substance, instead of a suspension.

Figures 4A-4D show in more detail, and on a larger scale, the filling head 28 used in the apparatus of Figure 3. The head comprises a substantially cylindrical body 40, the lower end of which is adapted, in use, to engage over the upper end of an aerosol container C. A tubular member 43 is mounted for slidable movement within the body 40. The tubular member 43 has a wide base portion 44, a narrower body portion 45 and a still narrower neck portion 46. In this context, "wide" and "narrow" refer to diameters.

The neck portion 46 of the tubular member 43 penetrates the base of an inverted cup 47, the wall of which surrounds the body portion 45. The body portion 45 can thus slide into the cup 47.

The base portion 44 of the tubular member abuts an inwardly extending lip 50 of the body 40. An outwardly extending lip 51 of the cup 47 rests on an internal shoulder 52 of the body 40.

A tubular pillar 55 is threadedly engaged with the body 40 such that its lower edge engages on the lip 51 of the cup 47. The pillar 55 thus fits around the wall of the cup 47.

A ring 56 is screwed into the pillar 55 so that it is mounted above the cup 47. The neck portion 46 of the tubular member 43 just enters into the ring 56. A sliding seal 58 is fitted between the ring 56 and the neck portion 46. Rubber O-ring seals 59, 60 are also provided between the ring 56 and the pillar 55 and in the base portion 44 of the member 43, where the nozzle of the container C fits (see later).

The ring 56 defines an upwardly tapering seat 62 for a correspondingly shaped plug 63. The plug 63 is biased into the seat by a compression spring 64, the upper end of which acts against an inwardly directed lip 65 of the pillar 55.

Above the lip 65 are opposed inlets 69,70 connecting from the exterior with the interior of the pillar 55, and thus with the interior of the tubular member 43, and so to the container.

Inlet pipes 72,73 are fitted into the inlets 69,70 respectively, sealed therein by O-ring seals 74,74'.

Between the inlets is provided a ball-bearing 75 which can close against either O-ring seal 74,74' to form a valve. As explained below, the ball bearing 75 is either forced against the O-ring 74 to seal the inlet 69 or against the O-ring 74' to seal the inlet 70.

Above the inlets 69,70, the pillar 55 provides a circular seat for a piston (not shown) which acts to press the fitting head down onto the container C.

The inlet 69 is connected to the line which carries the suspension from the metering cylinder 24 to the non-return valve 29. The inlet 70 is connected to the outlet of the metering cylinder 34 which contains propellant.

In its rest state, suspension S flows along the line from the metering cylinder 24 to the non-return valve 29 without entering the interior of the pillar 55, being prevented from doing so by the ball bearing valve 75, forced against the O-ring 74 by the over-pressure of propellant P in the line from cylinder 34. When it is desired to introduce a quantity of suspension into the container through the head 28, as described immediately below, the valves 23 and 29 are closed and the cylinder 26 operated to cause the piston therein to move downwardly.

At this stage the filling head 28 has been moved downwards, onto container C, as shown in Figure 4B. The nozzle of the container rests against O-ring seal 60 and, as the head is lowered, the nozzle forces the tubular member 43 into the cup 47 and ring 56 until the lip 50 of the body 40 abuts the rim of the container. In this position, the neck portion 46 of the tubular member 43 has penetrated through the ring 56 and pushes the plug 63 out of the seat 62, against the action of the spring 64. Communication between the interior of the pillar 55 and the interior of the tubular member 43 is now possible.

The increase in pressure of the suspension in the metering cylinder 24 which is caused by the operation of cylinder 26 is sufficient to overcome the force of the propellant on the ball bearing valve 75 and suspension is thus able to flow from the inlet 69, through the interior of the pillar 55 and the tubular member 43, and into the container. The inlet 70 remains closed since the pressure of the suspension forces the ball bearing valve 75 against O-ring 74'. Thus, suspension is unable to pass from the inlet 69 to the inlet 70 and contaminate the inlet 70.

The next stage in the filling of the container is to pass a quantity of propellant without any pharmaceutical material suspended therein into the head 28, through the interior of the pillar 55 and of the tubular member 43, and thence into the container. This is done by closing the valve 33 and operating the pneumatic cylinder 36. The increase in the pressure of the propellant which this causes is sufficient to move the ball bearing valve 75. Propellant is unable to pass from the now open inlet 70 to the inlet 69 because of the ball bearing valve 75 which is forced against O-

ring 74. This position is shown in Figure 4C.

Introducing propellant through the inlet 70 while the head 28 is still on container C flushes out suspension remaining in the interior of the pillar 55 and the tubular member 43. Accordingly, when the head 28 is lifted from the aerosol container after the filling operation has been completed, as shown in Figure 4D, such material as escapes from the lower end of the head consists substantially entirely of propellant, and no pharmaceutical material escapes into the surrounding atmosphere.

Figure 5 shows an alternative embodiment of a filling head 28 to be used in the apparatus of Figure 3. The head shown in Figure 5 comprises a substantially cylindrical body 80, the lower end of which is adapted, in use, to engage over the upper end of an aerosol container (not shown in this figure). A ring 81 is mounted for longitudinal sliding movement within the body 80 and has an inwardly directed flange 82 on which rests the lower end of a tubular member 83. The upper portion of the tubular member 83 is surrounded by an inverted cup 84. The cup 84 is in turn surrounded by the annular lower portion of a pillar 85. The annular portion is screw threaded into the body 80 to retain the cup 84 in place and sealing is provided by an O-ring seals 86 and a sliding seal 87. The tubular member 83 is urged into engagement with the flange 82 by a compression spring 88, the upper end of which bears against a face of the cup 84.

The pillar 85 has a pair of opposed inlets 89 and 90. The inlet 89 is connected to the line which carries suspension from the metering cylinder 24 to the non-return valve 29. The inlet 90 is connected to the outlet of the metering cylinder 34 which contains propellant. The inlets 89 and 90 communicate with the interior of the tubular member 83 via respective poppet valves 91 and 92 which are biased by compression springs 93 and 94 into their closed positions.

As in the embodiment of Figure 4, in its rest state, suspension flows along the line from the metering cylinder 24 to the non-return valve 29 without entering the interior of the tubular member 83, being prevented from doing so by the valve 91. When it is desired to introduce a quantity of suspension into the container through the head 28 the valves 23 and 29 are closed, and the cylinder 26 operated to cause the piston therein to move downwardly. The increase in pressure in the suspension in the metering cylinder 24 which is caused by this is sufficient to overcome the force of the spring 93 holding the valve 91 shut, and suspension is thus able to flow from the inlet 89, through the interior of the tubular member 83, into the container. The valve 92 remains closed, and indeed the effectiveness of the seal which it provides is increased by the head 95 of the valve 91 engaging the head 96 of the valve 92. Thus, suspension is unable to pass from the inlet 89 to the inlet 90 and contaminate the inlet 90.

The next stage in the filling of the container, as in the previous embodiment, is to pass a quantity of propellant without any pharmaceutical material suspended therein into the head 28, through the tubular member 83, and thence into the container. This is done by closing the valve 33 and operating the pneumatic cylinder 36. The increase in the pressure of the propellant which this causes is sufficient to open the valve 92 to permit propellant through the head 28. Propellant is unable to pass from the inlet 90 to the inlet 89 because of the action of the valve 91.

Introducing propellant through the inlet 90 while the head 28 is still on the container flushes suspension remaining in the interior of the tubular member 83 and in the space immediately above the tubular member. Accordingly, this embodiment also provides that when the head 28 is lifted from the aerosol container after the filling operation has been completed, such material as escapes from the lower end of the head consists substantially entirely of propellant, and no pharmaceutical material escapes.

Claims

1. A method of introducing into a container (c) a suspension or solution of a pharmaceutical substance in a propellant held under pressure, the suspension or solution being circulated in a line which includes a filling head (28), which method comprises bringing the filling head (28) into communication with the container (c); introducing a quantity of such suspension or solution into the container through the filling head (28); introducing a quantity of high pressure propellant without any of the said substance into the filling head (28) while it is still in communication with the container (c), thereby to flush through any suspension or solution remaining in the filling head (28); and withdrawing the filling head (28) from the container (c).
2. A method according to claim 1, wherein the propellant which contains the said substance and the propellant without any of the said substance are the same.
3. A method according to claim 1 or 2, wherein the propellant is 1,1,1,2-tetrafluoroethane.
4. A method according to any preceding claim, wherein the substance is salbutamol.
5. A method according to any of claims 1 to 3, wherein the substance is beclomethasone dipropionate.
6. An apparatus for introducing into a container (c) a suspension or solution of a pharmaceutical substance in a propellant held under pressure, which comprises a line in which the suspension or solution circulates, the line including a filling head (28) adapted to be brought into and out of communication with the container (c); means (21, 22, 23, 24) for supplying to the filling head (28) along the line a quantity of the said suspension or solution; and means (31, 32, 33, 34) for supplying to the filling head (28) a quantity of high pressure propellant without any of the said substance, the filling head (28) being so arranged that the flow of propellant without any of the said substance flushes out any suspension or solution remaining in the filling head (28).
7. An apparatus according to claim 6, wherein the filling head (28) comprises a passage for the suspension or solution and for the propellant alone, an inlet (70) for the suspension or solution and an inlet (69) for the propellant alone, each inlet (69,70) communicating with the said passage, and a valve member (75) movable between a position closing the inlet (70) for the suspension or solution and a position closing the inlet (69) for the propellant alone, whereby at any one time only one of the said inlets (69,70) is communicating with the said passage.
8. An apparatus according to claim 7, wherein the valve member is a spherical member (75).
9. An apparatus according to claim 6, wherein the filling head (28) comprises a passage for the suspension or solution and for the propellant alone, an inlet (70) for the suspension or solution and an inlet (69) for the propellant alone, each inlet (69,70) communicating with the said passage, and a valve member (95,96) in each inlet, the inlets (69,70) being opposite one another and the valve members (95,96) being arranged so that when one is open it presses the other further into its closed position.
10. A filling head for use in introducing into a container (c) a suspension or solution of a pharmaceutical substance in a propellant held under pressure, comprising an outlet adapted to communicate, in use, with the container (c); first (69) and second (70) inlets each communicating with the said outlet via a common flow path, through which inlets (69,70), in use, propellant containing the said substance, and high pressure propellant not containing the said substance, are respectively introduced; and means (75; 95,96) for selectively closing the first and second inlets (69,70) so that fluid entering through either inlet cannot flow out of the other.

11. A filling head according to claim 10, comprising a valve member (75) alternately closing the first and second inlets, the valve member being spherical member.

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12. A filling head according to claim 10, comprising a valve member (95,96) in each of the first and second inlets (69,70), the inlets being opposite one another and the valve member being so arranged that when one is open it presses the other further into its closed position.

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Patentansprüche

1. Verfahren zum Einführen einer Suspension oder Lösung einer pharmazeutischen Substanz in einem unter Druck gehaltenen Treibmittel in einen Behälter (c), wobei die Suspension oder Lösung in einer einen Füllkopf (28) aufweisenden Leitung zirkuliert, wobei das Verfahren die folgenden Schritte aufweist: Verbinden des Füllkopfes (28) mit dem Behälter (c); Einführen eines Quantums einer solchen Suspension oder Lösung durch den Füllkopf (28) in den Behälter; Einführen eines Quantums von Hochdrucktreibmittel ohne die genannte Substanz in den Füllkopf (28), während er noch mit dem Behälter (c) in Verbindung steht, wodurch jegliche in dem Füllkopf (28) verbleibende Suspension oder Lösung durchspült wird; und Abnehmen des Füllkopfes (28) von dem Behälter (c).

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2. Verfahren nach Anspruch 1, bei dem das die Substanz enthaltende Treibmittel und das Treibmittel ohne die genannte Substanz das gleiche sind.

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3. Verfahren nach Anspruch 1 oder 2, bei dem das Treibmittel 1,1,1,2-Tetrafluorethan ist.

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4. Verfahren nach einem der vorhergehenden Ansprüche, bei dem die Substanz Salbutamol ist.

5. Verfahren nach einem der Ansprüche 1 bis 3, bei dem die Substanz Beclomethason-Dipropionat ist.

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6. Vorrichtung zum Einführen einer Suspension oder Lösung einer pharmazeutischen Substanz in einem unter Druck gehaltenen Treibmittel in einen Behälter (c), mit einer Leitung, in der die Suspension oder Lösung zirkuliert, wobei die Leitung aufweist: einen Füllkopf (28), der derart ausgestaltet ist, daß er mit dem Behälter (c) in und außer Verbindung bringbar ist; Einrichtung (21,22,23,24) zum Zuführen eines Quantums der Suspension oder Lösung entlang der Leitung zu dem Füllkopf (28); und Einrichtungen

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(31,32,33,34) zum Zuführen in das Hochdrucktreibmittel Quantums ohne die genannte Substanz zu dem Füllkopf (28), wobei der Füllkopf (28) derart angeordnet ist, daß der Strom des Treibmittels ohne die genannte Substanz jede in dem Füllkopf (28) verbleibende Suspension oder Lösung hinausspült.

7. Vorrichtung nach Anspruch 6, bei der der Füllkopf (28) aufweist: einen Durchlaß für die Suspension oder Lösung und für das Treibmittel allein, einen Einlaß (70) für die Suspension oder Lösung und einen Einlaß (69) für das Treibmittel allein, wobei jeder Einlaß (69,70) mit dem Durchlaß in Verbindung steht, und ein Ventiltail (75), das zwischen einer Stellung zum Schließen des Einlasses (70) für die Suspension oder Lösung und einer Stellung zum Schließen des Einlasses (69) für das Treibmittel allein bewegbar ist, wodurch zu jeder Zeit nur einer der Einlässe (69,70) mit dem Durchlaß in Verbindung steht.

8. Vorrichtung nach Anspruch 7, bei der das Ventiltail ein sphärisches Teil (75) ist.

9. Vorrichtung nach Anspruch 6, bei der der Füllkopf (28) aufweist: einen Durchlaß für die Suspension oder Lösung und für das Treibmittel allein, einen Einlaß (70) für die Suspension oder Lösung und einen Einlaß (69) für das Treibmittel allein, wobei jeder Einlaß (69,70) mit dem Durchlaß in Verbindung steht, und ein Ventiltail (95,96) in jedem Einlaß, wobei die Einlässe (69,70) einander gegenüber liegen und die Ventiltail (95,96) derart angeordnet sind, daß, wenn eines geöffnet ist, es das andere weiter in seine geschlossene Stellung drückt.

10. Füllkopf zur Verwendung beim Einführen einer Suspension oder Lösung einer pharmazeutischen Substanz in einem unter Druck gehaltenen Treibmittel in einen Behälter (c), mit: einem Auslaß, der derart ausgestaltet ist, daß er im Gebrauch mit dem Behälter (c) in Verbindung steht; einem ersten (69) und zweiten (70) jeweils mit dem Auslaß über einen gemeinsamen Strömungsweg in Verbindung stehenden Einlaß, durch welche Einlässe (69,70) im Gebrauch die Substanz enthaltendes Treibmittel und Hochdrucktreibmittel ohne die genannte Substanz eingeführt werden; und Einrichtungen (75;95,96) zum selektiven Schließen des ersten und zweiten Einlasses (69,70) derart, daß in einen Einlaß eintretendes Fluid nicht aus dem anderen ausströmen kann.

11. Füllkopf nach Anspruch 10, der in Ventiltail (75) zum abwechselndem Schließen des ersten und

zweiten Einlässe aufweist, wobei das Ventiltail in sphärisches Teil ist.

12. Füllkopf nach Anspruch 10, der in j dem der ersten und zweiten Einlässe (69,70) ein Ventiltail (95,96) aufweist, wobei die Einlässe einander gegenüber liegen und die Ventiltile derart angeordnet sind, daß, wenn eines geöffnet ist, es das andere weiter in seine geschlossene Stellung drückt.

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Revendications

1. Méthode d'introduction dans un conteneur (c) d'une suspension ou d'une solution d'une substance pharmaceutique dans un propulseur sous pression, la suspension ou la solution circulant dans une conduite qui comporte une tête de remplissage (28), laquelle méthode comprend la mise en communication de la tête de remplissage (28) avec le conteneur (c); d'introduction d'une quantité d'une telle suspension ou solution dans le conteneur à travers la tête de remplissage (28); d'introduction d'une quantité de propulseur sous pression sans aucune desdites substances dans la tête de remplissage (28) alors qu'elle est encore en communication avec le conteneur (c), pour évacuer ainsi toute suspension ou solution restant dans la tête de remplissage (28); et le détachement de la tête de remplissage (28) du conteneur (c).
2. Méthode selon la revendication 1, dans laquelle le propulseur qui contient ladite substance et ledit propulseur sans aucune desdites substances sont les mêmes.
3. Méthode selon la revendication 1 ou 2, dans laquelle le propulseur est du 1,1,1,2-tétrafluoroéthane.
4. Méthode selon l'une quelconque des revendications précédentes, dans laquelle la substance est du salbutamol
5. Méthode selon l'une quelconque des revendications 1 à 3, dans laquelle la substance est du dipropionate de beclométhasone.
6. Appareil pour introduire dans un conteneur (c) une suspension ou une solution d'une substance pharmaceutique dans un propulseur sous pression, qui comprend une conduite dans laquelle circule la suspension ou la solution, la ligne comportant un tête de remplissage (28) adaptée pour être mise en et hors communication avec le conteneur (c); un moyen (21, 22, 23, 24) pour

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fournir à la tête de remplissage (28) par la conduite, une quantité de ladite suspension ou solution, et un moyen (31, 32, 33, 34) pour fournir à la tête de remplissage (28) une quantité de propulseur sous haute pression sans aucune desdites substances, la tête de remplissage (28) étant agencée de telle sorte que le flux de propulseur sans aucune desdites substances évacue toute suspension ou solution restant dans la tête de remplissage (28).

7. Appareil selon la revendication 6, dans lequel la tête de remplissage (28) comprend un passage pour la suspension ou la solution et pour le propulseur seul, une entrée (70) pour la suspension ou solution et une entrée (69) pour le propulseur seul, chaque entrée (69, 70) communiquant avec ledit passage, et une pièce formant valve (75) mobile entre une position fermant l'entrée (70) pour la suspension ou la solution et une position fermant l'entrée (69) pour le propulseur seul, à tout moment une seule desdites entrées (69, 70) est ainsi en communication avec ledit passage.
8. Appareil selon la revendication 7, dans lequel la pièce formant valve est une pièce sphérique (75).
9. Appareil selon la revendication 6, dans lequel la tête de remplissage (28) comprend un passage pour la suspension ou la solution et pour le propulseur seul, une entrée (70) pour la suspension ou la solution et une entrée (69) pour le propulseur seul, chaque entrée (69, 70) communiquant avec ledit passage, et une pièce formant valve (95, 96) dans chaque entrée, les entrées (69, 70) étant opposées l'une à l'autre et les pièces formant valve (95, 96) étant agencées de sorte que lorsque l'une est ouverte elle appuie davantage l'autre dans sa position fermée.
10. Tête de remplissage à utiliser pour l'introduction dans un conteneur (c) d'une suspension ou d'une solution d'une substance pharmaceutique dans un propulseur sous pression, comprenant une sortie adaptée pour communiquer, lors de l'utilisation avec le conteneur (c); des première (69) et seconde (70) entrées communiquant avec ladite sortie via un chemin de flux commun, à travers lesquels entrées (69, 70) on introduit, lors de l'utilisation, respectivement un propulseur contenant ladite substance, et un propulseur sous haute pression ne contenant pas ladite substance; et un moyen (75; 95, 96) pour fermer sélectivement les première et seconde entrées (69, 70) afin que le fluide entrant par n'importe quelle entrée ne puisse s'écouler par l'autre.
11. Tête de remplissage selon la revendication 10,

compr nant une pièc formant valv (75) fermant alternativement les pr mière et seconde entrées, la pièce formant valve étant une pièce sphérique.

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12. Tête de remplissage selon la revendication 10, comprenant une pièce formant valve (95, 96) dans chacune des première et seconde entrées (69, 70), les entrées étant opposées l'une à l'autre et la pièce formant valve étant agencée de telle sorte que lorsque l'une est ouverte elle appuie davantage l'autre dans sa position fermée.

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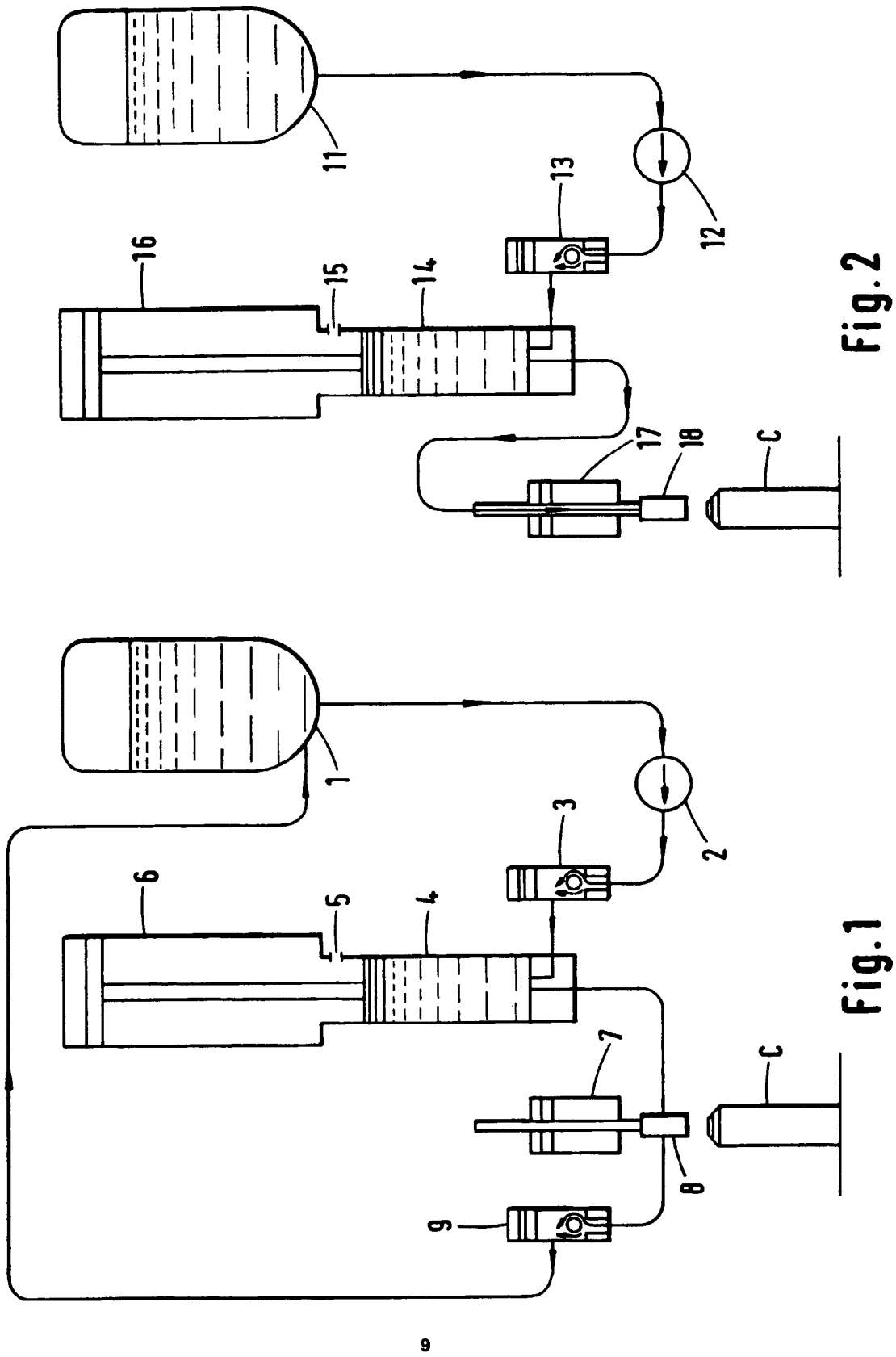
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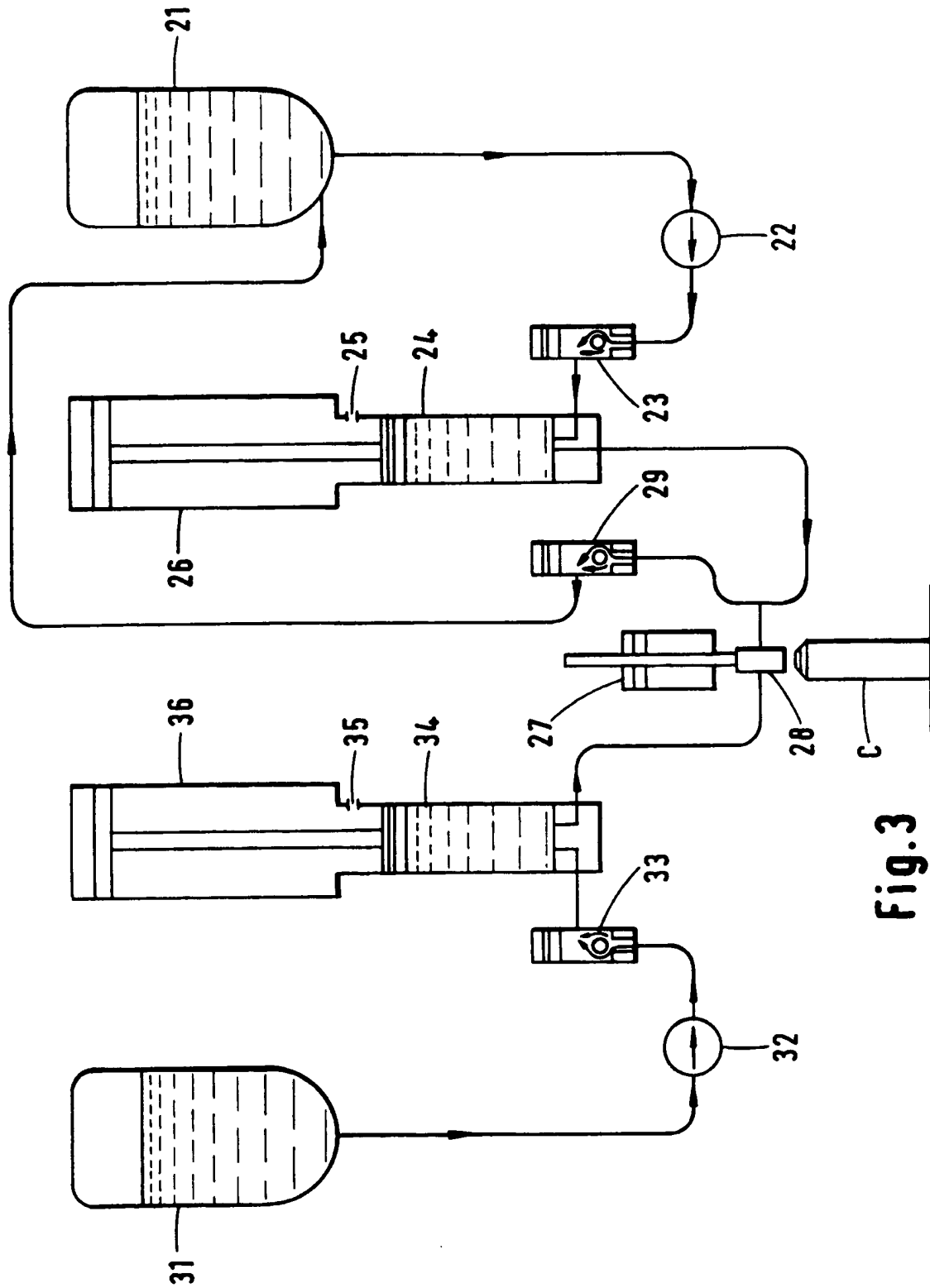


Fig.3

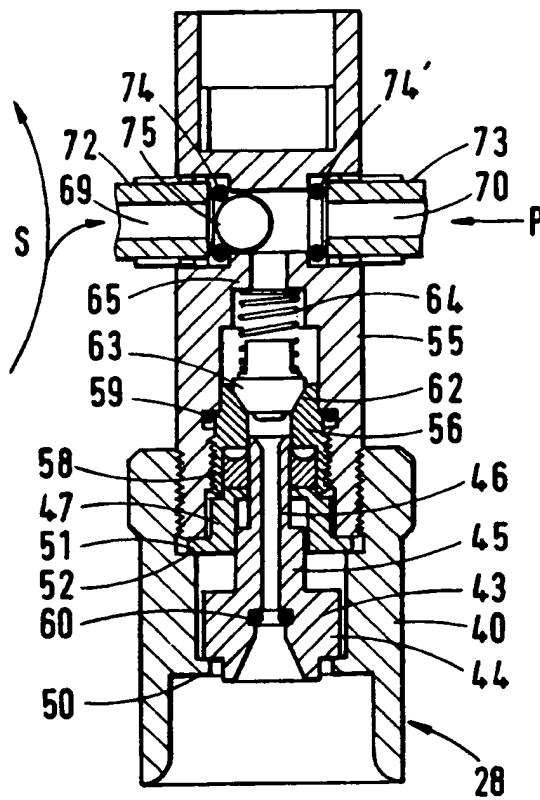


Fig. 4(a)

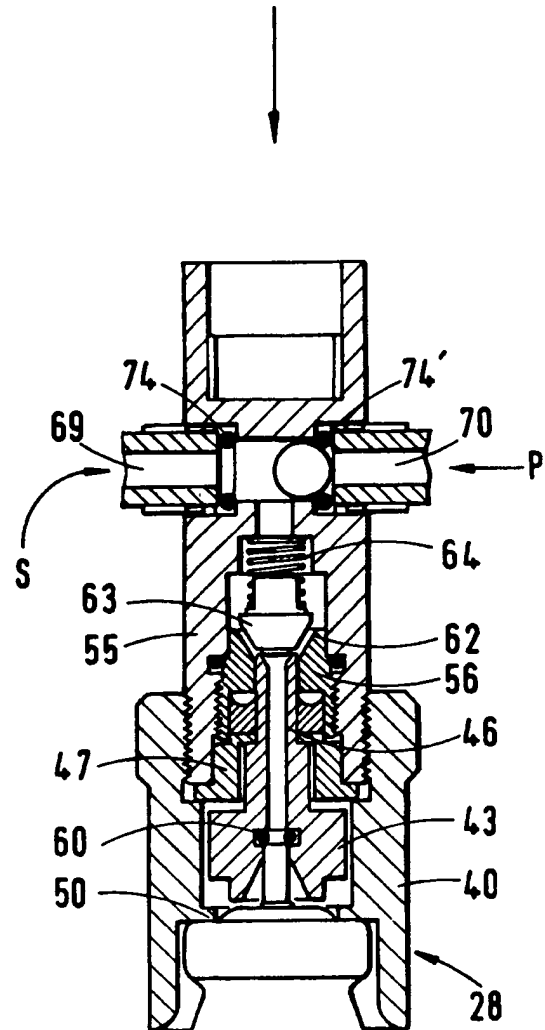


Fig. 4(b)

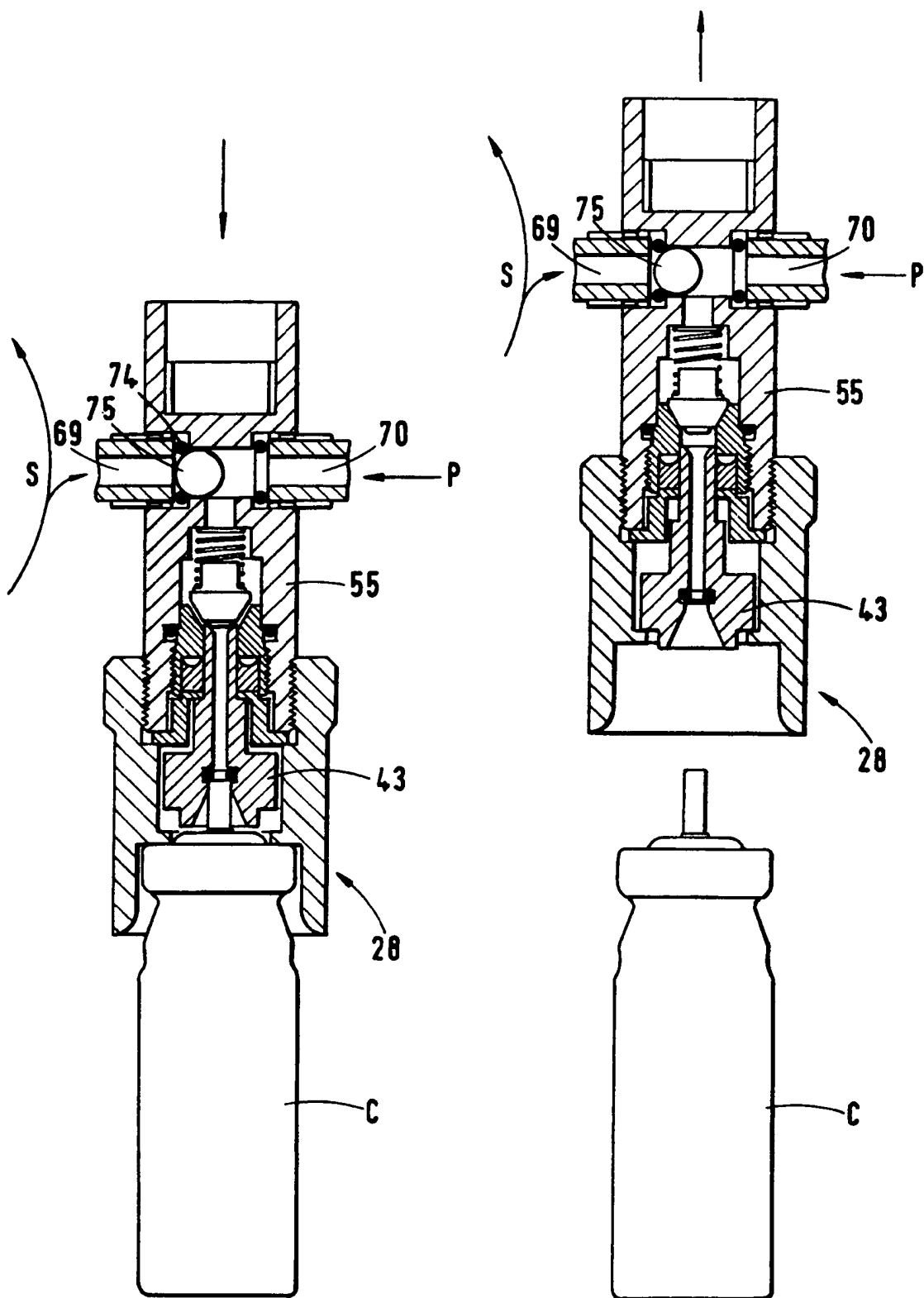


Fig.4(c)

Fig.4(d)

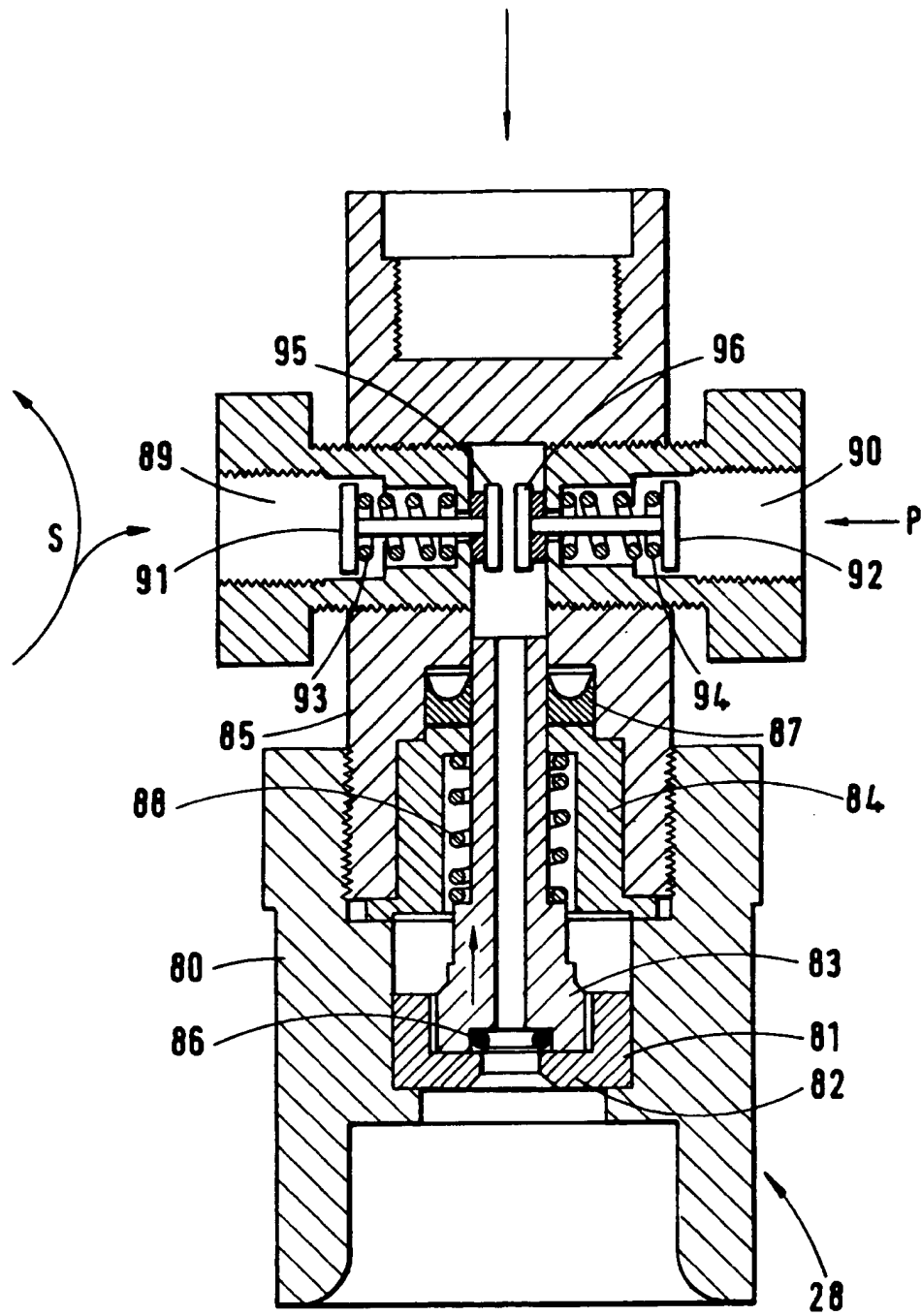


Fig.5